|  |  |                                | Sheet 1 (A) of 12             |
|--|--|--------------------------------|-------------------------------|
| SUBSTITUTE FORM PTO-1449<br>(MODIFIED) | U.S. DEPARTMENT OF COMMERCE<br>PATENT AND TRADEMARK OFFICE | ATTY. DOCKET NO.<br>RENAS/04   | SERIAL NO.<br>09/939,962 OIPE |
|  | ON DISCLOSURE<br>Γ BY APPLICANT                            | APPLICANT<br>Michael B. Foster | APR 0.5 ->e/                  |
| (Use several s<br>(37 CFR 1.98(b))     | sheets if necessary)                                       | FILING DATE<br>August 27, 2001 | GROUP 1631 6                  |
|  | U.S. PATENT DOCUMENTS                                      |                                | PADEMARK                      |
|  |  | T T                            |                               |

| (37 CFR 1.98(                         | b)) |            |                |                |                       |       |       |                 |                                   |            |   | August 2   | 27, 2001                  | -1635  | 162                                     |
|---------------------------------------|-----|------------|----------------|----------------|-----------------------|-------|-------|-----------------|-----------------------------------|------------|---|------------|---------------------------|--|---|
|                                       |     |            |                |                |                       |       |       |                 | U.S.                              | PATENT     | DOCUMENTS                                 |            |                           |  | PADEMAN                                 |
| EXAMINER<br>INITIAL                   |     | _          | PA             | TEN            | IT N                  | IUM   | BEF   | ₹               | ISSUE PATENTEE                    |            |   |            | CLASS                     | SUBCLASS   | FILING DATE<br>IF APPROPRIAT            |
| CPZ.                                  | A.A | 5          | 8              | 5              | 5                     | 9     | 2     | 0               | 01/1999                           | Chien      |   | 1          |                           |  |   |
|                                       | A.B |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.C |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.D |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.E |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.F |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.G |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.H |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | I.A |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.J |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
| · · · · · · · · · · · · · · · · · · · | A.K | T          |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       |     | •          | 1              |                |                       | FOR   | EIG   | N PA            | TENTS OR PU                       | JBLISHE    | D FOREIGN PAT                             | ENT APP    | PLICATION                 | 5  |   |
|                                       |     |            |                |                |                       |       |       | P               | UBLICATION<br>DATE                |            | COUNTRY OR<br>PATENT OFFIC                |            | CLASS                     | SUBCLASS   | TRANSLATION<br>(YES/NO)                 |
|                                       | A.L |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.M |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.N |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.O |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  |   |
|                                       | A.P |            |                | -              |                       |       |       |                 |                                   |            |   |            |                           | 9  |   |
|                                       | A.Q |            |                |                |                       |       |       |                 |                                   |            |   |            |                           |  | <u></u>                                 |
|                                       | -   |            |                |                |                       | ОТ    | HEF   | l DO            | CUMENTS (Inc                      | cluding A  | uthor, Title, Date,                       | Place of   | Publication)              |  |   |
| CPL                                   | A.R |            | entii<br>98, 6 |                |                       | am a  | ınd a | bstra           | cts, <u>Consultants</u>           | s Confere  | nce on Adult Grov                         | vth Horm   | one Deficien              | cy Genentech, Inc.,                              | San Francisco, CA,                      |
|                                       | A.S | An<br>afte | nato<br>er Gl  | et al<br>H rej | ., <u>Bo</u><br>place | ody o | om    | oositi<br>erapy | on, bone metab<br>at low doses Jo | olism, and | d heart structure a<br>Clinical Endocrine | nd functio | n in growth<br>Metabolism | h <u>ormone (GH)-def</u><br>, Vol. 77, No. 6, 19 | cient adults before a<br>93, 1671-1676. |
|                                       | A.T | Αn         | drew           | /s, <u>F</u>   | orev                  | er y  | ouns  | , Sma           | art Money, 2000                   | 0, 127-132 | 2.  |            |                           |  |   |

EXAMINER

DATE CONSIDERED 6/14/04

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and to considered. Include a copy of this form with next communication to applicant.

Sheet 2 (B) of SERIAL NO. ATTY. DOCKET NO. U.S. DEPARTMENT OF COMMERCE **SUBSTITUTE FORM PTO-1449** 09/939,962 PATENT AND TRADEMARK OFFICE RENAS/04 (MODIFIED) APPLICANT INFORMATION DISCLOSURE STATEMENT BY APPLICANT Michael B. Foster GROUN **FILING DATE** (Use several sheets if necessary) August 27, 2001 1055 (37 CFR 1.98(b)) U.S. PATENT DOCUMENTS **FILING DATE ISSUE EXAMINER** IF APPROPRIATE **PATENTEE CLASS SUBCLASS** PATENT NUMBER DATE INITIAL B.A B.BB.C B.D B.E B.F B.G B.H B.I B.JB.K FOREIGN PATENTS OR PUBLISHED FOREIGN PATENT APPLICATIONS TRANSLATION DOCUMENT **PUBLICATION COUNTRY OR CLASS SUBCLASS** (YES/NO) PATENT OFFICE DATE NUMBER B.L B.MB.N B.O B.P B.Q OTHER DOCUMENTS (Including Author, Title, Date, Place of Publication) a Attanasio et al., Adult growth hormone (GH)-deficient patients demonstrate heterogeneity between childhood onset and adult onset B.R before and during human GH treatment, Journal of Clinical Endocrinology and Metabolism, Vol. 82, No. 1, 1997, 82-88. Bates et al., The effect of hypopituitarism on life expectancy Journal of Clinical Endocrinology and Metabolism, Vol. 81, No. 3, 1996, B.S 1169-1172. Bengtsson, Efficacy of growth hormone-replacement therapy: quality of life and performance The Endocrinologist, Vol. 8, No. 6,

**EXAMINER** 

Suppl Lippincott Williams & Wilkins, 1998, 31S-35S.

B.T

DATE CONSIDERED

6/14/04

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance anchot considered. Include a copy of this form with next communication to applicant.

Sheet 3 (C) of 12

Substitute Disclosure Form (PTO-1449)

| SUBSTITUTE FORM PTO-1449 U.S. DEPARTMEN' (MODIFIED) PATENT AND TRA |                |              |       |  |                 |             |  |                      |                              | ATTY<br>RENA | '. DOCKET NO<br>AS/04  | D.<br>      | 09/939,962 \ P E |                               |  |  |
|--|----------------|--------------|-------|--|-----------------|-------------|--|----------------------|------------------------------|--------------|------------------------|-------------|------------------|-------------------------------|--|--|
|  |                |              |       |  |                 |             | LOSURE<br>LICANT                       |                      |                              |              | ICANT<br>sel B. Foster |             |                  | 2 5 3004                      |  |  |
|  |                |              | (Use  | sevei  | al sheet        | ts if no    | ecessary)                              |                      |                              |              | G DATE                 | -           | GROU<br>1655     | APR 0 5 2000 W                |  |  |
| (37 CFR 1.98(  | b))            |              |       | <del></del>  |                 |             |  |                      |                              | A CALLERY    |                        |             |                  |                               |  |  |
|  | <del></del>    |              |       |  |                 |             | U.S. P                                 | ATENT                | DOCUMENTS                    |              | <del></del>            | 1           |                  | PADE                          |  |  |
| EXAMINER<br>INITIAL  |                | ,            | PATE  | NT N   | UMBE            | ISSUE PATEN |  |                      |                              | ·            | CLASS                  | SUBC        | CLASS            | FILING DATE<br>IF APPROPRIATE |  |  |
|  | C.A            |              |       |  |                 | ↓           |  |                      |                              | ···········  | ļ <u> </u>             | ļ           |                  |                               |  |  |
|  | C.B            |              |       | $oldsymbol{ol}}}}}}}}}}}}}}}}}}$ |                 |             |  |                      |                              |              |                        | ļ           |                  |                               |  |  |
|  | C.C            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.D            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.E            |              |       |  |                 |             |  |                      |                              | ,            |                        | <u></u>     |                  |                               |  |  |
|  | C.F            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.G            | $\prod$      |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.H            |              | 1     |  |                 |             |  | -                    |                              |              |                        |             |                  |                               |  |  |
|  | C.I            | 11           |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.J            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.K            | 11           |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  |                |              |       | ]  | FOREIC          | GN PA       | ATENTS OR PU                           | BLISHE               | O FOREIGN PA                 | TENT A       | PPLICATION             | S           |                  |                               |  |  |
|  |                | J            |       | UMEI<br>MBEI   |                 | F           | PUBLICATION DATE                       |                      | COUNTRY OF PATENT OFFICE     |              | CLASS                  | SUBC        | LASS             | TRANSLATION<br>(YES/NO)       |  |  |
|  | C.L            |              |       | -  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.M            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.N            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.O            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
| •  | C.P            |              |       |  | -               |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | C.Q            |              |       |  |                 |             |  |                      |                              |              |                        |             |                  |                               |  |  |
|  | · <del>-</del> | •            |       |  | ОТНЕ            | R DO        | CUMENTS (Inc                           | luding A             | uthor, Title, Date           | , Place      | of Publication)        |             |                  |                               |  |  |
| an   | C.R            | Beng<br>Endo | tssor | et al.   | , <u>Treatn</u> | nent o      | of adults with gro<br>sm, Vol. 76, No. | wth horn<br>2, 1993, | none (GH) defici<br>309-317. | ency wit     | th recombinant         | human Gl    | J Journal        | of Clinical                   |  |  |
|  | C.S            | Beng         | tssor | , The  | conseq          | uence       | s of growth horn                       | one defic            | ciency in adults             | Acta End     | locrinologica, \       | √o1. 128, S | Supp 2, 19       | 993, 2-5.                     |  |  |
|  | C.T            |              |       | icacy<br>5S-21   |                 | th ho       | rmone-replaceme                        | ent therap           | y: body compos               | sition an    | d bone density         | The Endo    | crinologis       | t, Vol. 8, No. 6, Supp        |  |  |
| EXAMINER   |                | 26           |       | 1  |                 | /           |  |                      | DATE CONSI                   | DEREC        | 6/1                    | 4/0         | 4                |                               |  |  |
| EXAMINER:  |                |              |       |  |                 |             |  | ce. Draw             | line through cit             | ation on     | ly if not in con       | formance_   | andhot co        | nsidered. Include a           |  |  |

Sheet 4 (D) of 12 ATTY, DOCKET NO. SERIAL NO. U.S. DEPARTMENT OF COMMERCE SUBSTITUTE FORM PTO-1449 09/939,962 PATENT AND TRADEMARK OFFICE RENAS/04 (MODIFIED) APPLICANT INFORMATION DISCLOSURE Michael B. Foster STATEMENT BY APPLICANT FILING DATE GRd (Use several sheets if necessary) August 27, 2001 (37 CFR 1.98(b)) U.S. PATENT DOCUMENTS FILING DATE **ISSUE EXAMINER** SUBCLASS IF APPROPRIATE **CLASS PATENTEE** PATENT NUMBER DATE INITIAL D.A D.BD.C D.D D.E D.F D.G D.H D.I D.J D.K FOREIGN PATENTS OR PUBLISHED FOREIGN PATENT APPLICATIONS TRANSLATION **PUBLICATION COUNTRY OR DOCUMENT** (YES/NO) **SUBCLASS** PATENT OFFICE **CLASS** NUMBER DATE D.L D.M D.N D.O D.P D.O OTHER DOCUMENTS (Including Author, Title, Date, Place of Publication) Blethen, Dosing, monitoring, and safety of growth hormone-replacement therapy in adults with growth hormone deficiency The D.R Endocrinologist, Vol. 8, No. 6, Supp 1,1998, 36S-40S. Brody, Restoring ebbing hormones may slow aging The New York Times, July 18, 1995, late edition, Sec. C, Page 1, Col. 1, 2-10. D.S

DATE CONSIDERED 6/14/04

D.T

Medicine, Vol. 11, No. 4, 1995, 735-748.

Carter, Effect of anabolic hormones and insulin-like growth factor-l on muscle mass and strength in elderly persons Clinics in Geriatric

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and to considered. Include a copy of this form with next communication to applicant.

Sheet 5 (E) of 12

| SUBSTITUTE (MODIFIED) | FORM P      | ГО-1449           |                  |                          |                      | U.S. DEPARTM<br>PATENT AND T | ENT OF COMMERCE TRADEMARK OFFICE | ATTY. DOCKET NO. S<br>RENAS/04 0      |                      |             |               | SERIAL NO. 0 1 P E 10 19/939,962 |  |  |
|-----------------------|-------------|-------------------|------------------|--------------------------|----------------------|------------------------------|----------------------------------|---------------------------------------|----------------------|-------------|---------------|----------------------------------|--|--|
|                       |             |                   |                  |                          |                      | LOSURE<br>PLICANT            |                                  |                                       | CANT<br>el B. Foster |             |               | APR 0 5 2004                     |  |  |
|                       |             | J)                | Jse sev          | eral sh                  | eets if n            | ecessary)                    |                                  |                                       | G DATE               |             | GROU          | 1631                             |  |  |
| (37 CFR 1.98(b        | ))          |                   |                  |                          |                      |                              |                                  | Augus                                 | t 27, 2001           |             | 1653-         | MADEMARK                         |  |  |
|                       |             |                   |                  |                          |                      | U.S. P                       | ATENT DOCUMENTS                  |                                       | <del></del>          | <u></u>     |               |                                  |  |  |
| EXAMINER<br>INITIAL   |             | PA                | TENT             | NUM                      | BER                  | ISSUE<br>DATE                | PATENTEE                         |                                       | CLASS                | SUBC        | LASS          | FILING DATE<br>IF APPROPRIATE    |  |  |
|                       | E.A         |                   |                  |                          |                      |                              |                                  |                                       |                      | ļ <u> </u>  |               |                                  |  |  |
|                       | E.B         |                   |                  | Ш                        |                      |                              |                                  |                                       |                      | <u> </u>    |               |                                  |  |  |
|                       | E.C         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.D         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.E         |                   |                  |                          |                      |                              |                                  |                                       |                      | <u> </u>    |               |                                  |  |  |
|                       | E.F         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.G         |                   |                  |                          |                      |                              |                                  |                                       | ļ                    |             |               |                                  |  |  |
|                       | E.H         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.I         |                   |                  |                          |                      |                              | ·                                |                                       |                      |             |               |                                  |  |  |
|                       | E.J         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.K         |                   |                  |                          |                      |                              |                                  |                                       | <u> </u>             | <u> </u>    | n · · · · · · |                                  |  |  |
|                       |             |                   |                  | FOR                      | EIGN PA              | ATENTS OR PU                 | BLISHED FOREIGN PA               | TENT A                                | PPLICATION           | s           |               |                                  |  |  |
|                       |             |                   | OCUM<br>NUMB     |                          | ]                    | PUBLICATION<br>DATE          | COUNTRY OF                       |                                       | CLASS                | SUBCI       | LASS          | TRANSLATION<br>(YES/NO)          |  |  |
|                       | E.L         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.M         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.N         |                   |                  |                          |                      |                              | 1.7                              |                                       |                      |             |               |                                  |  |  |
|                       | E.O         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.P         |                   |                  |                          |                      |                              |                                  |                                       |                      |             |               |                                  |  |  |
|                       | E.Q         |                   |                  |                          |                      |                              |                                  | · · · · · · · · · · · · · · · · · · · |                      |             |               |                                  |  |  |
|                       | •           | _                 |                  | ОТ                       | HER DO               | CUMENTS (Inc                 | luding Author, Title, Date       | e, Place o                            | of Publication)      |             |               |                                  |  |  |
| cor                   | E.R         | Cuneo,<br>1998, 2 | Effica<br>228-30 | icy of s<br>S.           | growth h             | ormone-replacem              | ent therapy: lipid and ca        | rdiovasc                              | ular effects, Th     | e Endocrir  | nologist,     | Vol. 8, No. 6, Supp 1,           |  |  |
|                       | E.\$        | Cuneo<br>Vol. 42  | et al.,          | <u>Growth</u><br>12, 199 | n hormor<br>13, 1519 | ne treatment impr<br>-1523.  | oves serum lipids and lip        | oproteins                             | s in adults with     | growth ho   | ormone d      | eficiencyMetabolism,             |  |  |
|                       | E.T         | de Boe            | r et al.         | , <u>Serun</u>           | n lipid le           | vels in growth ho            | ormone-deficient men Me          | tabolism                              | , Vol. 43, No. 2     | 2, 1994, 19 | 99-203.       |                                  |  |  |
| EXAMINER              | 0           |                   | 11               | 1                        | /                    |                              | DATE CONS.                       | IDERED                                | 6/19                 | 4/04        | _             |                                  |  |  |
| EXAMINER: I           | nitial if c | itation co        | onsider          | ed, wh                   | ether or             | not in conforman             | ce. Draw line through cit        | tation on                             | ly if not in con     | formance:   | anchot co     | nsidered. Include a              |  |  |
| copy of diffs for     | 7101 11     | -71 00110         |                  |                          | -pp.iom              | ·····                        |                                  |                                       |                      | Substit     | ute Discl     | osure Form (PTO-1449)            |  |  |

| SUBSTITUTE I                 | FORM PT      | O-1449           |                 |                        |                               | U.<br>PA            | S. DEPARTI<br>TENT AND      | MENT OF (                   | COMMERCE<br>ARK OFFICE             | ATTY.<br>RENAS                               | DOCKET NO<br>5/04   | ,           | SERIAL NO.<br>09/939,962 |                               |  |
|------------------------------|--------------|------------------|-----------------|------------------------|-------------------------------|---------------------|-----------------------------|-----------------------------|------------------------------------|--|---------------------|-------------|--------------------------|-------------------------------|--|
|                              |              | IN<br>ST         | FORN<br>ATE     | AAT<br>MEN             | ION DI                        | SCLO<br>APPLI       | SURE<br>CANT                |                             | !                                  | APPLIO<br>Michae                             | CANT<br>I B. Foster |             |                          | 1                             |  |
| (37 CFR 1.98(b               | ))           | J)               | lse sev         | /eral                  | sheets i                      | if nece             | ssary)                      |                             |                                    | FILING DATE August 27, 2001 GROUP APR 0 5 20 |                     |             |                          |                               |  |
|                              |              |                  |                 |                        |                               |                     | U.S.                        | PATENT I                    | OCUMENTS                           |  |                     |             |                          | TA MADEWARK OF                |  |
| EXAMINER<br>INITIAL          |              | PA               | TENT            | ' אע                   | MBER                          |                     | ISSUE<br>DATE               |                             | PATENTEE                           |  | CLASS               | SUBCLASS    |                          | FILING DATE<br>IF APPROPRIATE |  |
|                              | F.A          |                  |                 |                        |                               |                     |                             |                             |                                    |  | ļ                   |             |                          |                               |  |
|                              | F.B          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.C          |                  |                 | $\perp$                |                               |                     |                             | <u> </u>                    |                                    |  | ļ <u>.</u>          |             |                          |                               |  |
|                              | F.D          |                  |                 |                        |                               |                     |                             | <u>.</u>                    |                                    |  | _                   |             |                          |                               |  |
|                              | F.E          |                  |                 |                        |                               |                     | _                           |                             |                                    |  | <u> </u>            |             |                          |                               |  |
|                              | F.F          |                  |                 |                        | $\perp \downarrow \downarrow$ |                     |                             | ļ                           |                                    |  | <u> </u>            |             | <del>"</del>             |                               |  |
|                              | F.G          |                  | Ш               |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.H          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.I          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.J          |                  |                 |                        |                               |                     |                             |                             |                                    | <u></u>                                      |                     |             |                          |                               |  |
|                              | F.K          |                  | Ш               |                        |                               |                     |                             | <u> </u>                    |                                    |  | <u> </u>            |             |                          |                               |  |
| ii <u>.</u>                  |              |                  |                 | FC                     | OREIGN                        | TA9                 | ENTS OR P                   | UBLISHED                    | FOREIGN PA                         | TENT A                                       | PPLICATION          | <u> </u>    |                          |                               |  |
|                              |              |                  | CUM             |                        | r                             | PUBLICATION<br>DATE |                             |                             | COUNTRY O                          |  | CLASS               | SUBCI       | LASS                     | TRANSLATION<br>(YES/NO)       |  |
|                              | F.L          |                  |                 |                        |                               |                     |                             |                             |                                    | į,   | 4                   |             |                          |                               |  |
|                              | F.M          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.N          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             | - 1                      |                               |  |
|                              | F.O          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.P          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              | F.Q          |                  |                 |                        |                               |                     |                             |                             |                                    |  |                     |             |                          |                               |  |
|                              |              |                  |                 |                        |                               |                     |                             |                             | thor, Title, Da                    |  |                     |             |                          |                               |  |
| CK                           | F.R          | Eden e           | t al., <u>C</u> | <u>irow</u><br>ns, A   | th horm<br>rteriosc           | one tr              | eatment of g                | rowth horm<br>bosis, Vol.   | one-deficient a<br>13, No. 2, 1993 | <u>dults resu</u><br>, 296-301               | lts in a marked     | l increase_ | in Lp(a) a               | and HDL cholesterol           |  |
|                              | F.S          |                  |                 |                        |                               |                     |                             |                             | Res, Vol. 40, 19                   |  |                     |             |                          |                               |  |
|                              | F.T          | Hoffm<br>Clinica | an et a         | al., <u>A</u><br>ocrir | dults w                       | ith gro             | owth hormon<br>etabolism, V | e deficiency<br>ol. 80, No. | y have abnorma<br>1, 1995, 72-77.  | l body co                                    | mposition but       | normal en   | ergy meta                | abolismJournal of             |  |
| EXAMINER                     | <i>Q</i>     | De               | /               | 1                      | /                             |                     |                             |                             | DATE CONS                          |  | 6/19                |             |                          |                               |  |
| EXAMINER:<br>copy of this fo | Initial if c | itation com      | onside<br>munic | red,<br>ation          | whether                       | r or no             | t in conform                | ance. Draw                  | line through c                     | itation on                                   | ly if not in cor    |             |                          | nsidered. Include a           |  |

Substitute Disclosure Form (PTO-1449)

Sheet 6 (F) of 12

Sheet 8 (H) of 12 ATTY. DOCKET NO. SERIAL NO. U.S. DEPARTMENT OF COMMERCE SUBSTITUTE FORM PTO-1449 09/939,962 PATENT AND TRADEMARK OFFICE RENAS/04 (MODIFIED) APPLICANT INFORMATION DISCLOSURE Michael B. Foster STATEMENT BY APPLICANT **GROUP FILING DATE** (Use several sheets if necessary) August 27, 2001 d653 (37 CFR 1.98(b)) U.S. PATENT DOCUMENTS **FILING DATE ISSUE EXAMINER SUBCLASS** IF APPROPRIATE **CLASS PATENTEE** PATENT NUMBER DATE INITIAL H.A H.B H.C H.D H.E H.F H.G H.H H.I H.J H.K FOREIGN PATENTS OR PUBLISHED FOREIGN PATENT APPLICATIONS TRANSLATION COUNTRY OR **PUBLICATION DOCUMENT SUBCLASS** (YES/NO) **CLASS** DATE **PATENT OFFICE** NUMBER H.L H.M H.N H.O H.P H.Q OTHER DOCUMENTS (Including Author, Title, Date, Place of Publication) Kleinberg and Melmed, The adult growth hormone deficiency syndrome: signs, symptoms, and diagnosis The Endocrinologist, Vol. 8, H.R No. 6, Supp 1, 1998, 8S-14S. Lieberman and Hoffman, Growth hormone deficiency in adults: characteristics and response to growth hormone replacements. H.S Pediatrics, Vol. 128, No. 5, Part 2, 1996, S58-S60. Meling and Nylen, Growth hormone deficiency in adults: a review The American Journal of the Medical Sciences, Vol. 311, No. 4,

**EXAMINER** 

H.T

1996, 153-166.

DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and to considered. Include a copy of this form with next communication to applicant.

| SUBSTITUTE<br>(MODIFIED) | FORM PT      | O-1449            |                    |                               |                       |   | ENT OF COMMERCE<br>RADEMARK OFFICE               |                             | ATTY. DOCKET NO. RENAS/04 SERIAL NO. 09/939,962 |                          |                         |            |                       |
|--------------------------|--------------|-------------------|--------------------|-------------------------------|-----------------------|---|--|-----------------------------|---|--------------------------|-------------------------|------------|-----------------------|
| ,                        |              |                   |                    |                               |                       | LOSURE<br>LICANT                        |  | APPLIC<br>Michael           | ANT<br>B. Foster                                |                          |                         |            | - ~ - 2004            |
| (37 CFR 1.98(E           | o))          | J)                | Jse s <b>ev</b>    | eral she                      | ets if ne             | ecessary)                               |  | FILING<br>August            |   |                          | GROUI                   | A A        | PR 0 5 2004<br>1631   |
| (5. 5                    |              |                   |                    |                               |                       | U.S. P                                  | ATENT DOCUMENTS                                  |                             |   |                          |                         | N.         | TRADEMARY             |
| EXAMINER<br>INITIAL      |              | PA                | TENT               | NUMB                          | ER                    | ISSUE<br>DATE                           | PATENTEE   |                             | CLASS   | SUBCLASS                 |                         | FIL        | ING DATE<br>PROPRIATE |
|                          | G.A          |                   |                    | $\prod$                       |                       |   |  |                             |   |                          |                         |            |                       |
| <del></del>              | G.B          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.C          |                   |                    |                               |                       |   |  |                             |   |                          | ***                     |            |                       |
|                          | G.D          |                   |                    |                               |                       |   |  |                             |   |                          | <u>.</u>                |            | <u>.</u>              |
|                          | G.E          |                   |                    |                               |                       |   |  |                             |   |                          |                         | <u> </u>   | <del></del>           |
|                          | G.F          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.G          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.H          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.I          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.J          |                   |                    |                               |                       |   |  | ·                           | _   |                          |                         | ļ <u> </u> |                       |
|                          | G.K          |                   |                    |                               |                       |   |  |                             |   |                          |                         | <u> </u>   |                       |
|                          |              |                   |                    | FORE                          | IGN PA                | ATENTS OR PU                            | BLISHED FOREIGN PA                               | TENT AP                     | PLICATION                                       | S                        | ···· -                  |            |                       |
|                          |              |                   | OCUM<br>NUMB       |                               | · F                   | PUBLICATION<br>DATE                     | COUNTRY O<br>PATENT OFFI                         |                             | CLASS   | SUBCI                    | LASS                    |            | NSLATION<br>YES/NO)   |
|                          | G.L          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.M          |                   |                    |                               |                       |   |  |                             |   |                          | · · ·                   |            |                       |
|                          | G.N          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.0          |                   |                    |                               |                       |   |  |                             |   | •                        |                         |            |                       |
|                          | G.P          |                   |                    |                               |                       |   |  |                             |   |                          |                         |            |                       |
|                          | G.Q          |                   |                    |                               |                       |   | <u> </u>   |                             |   |                          |                         |            |                       |
|                          |              |                   |                    | OTH                           | ER DC                 | CUMENTS (Incl                           | luding Author, Title, Da                         | te, Place of                | Publication)                                    |                          |                         |            |                       |
| cor                      | G.R          | Hollov<br>elderly | vay et a           | al., <u>Effec</u><br>n, Journ | ets of re<br>al of Cl | combinant huma<br>inical Endocrinol     | n growth hormone on me<br>logy and Metabolism, V | etabolic inc<br>ol. 72, No. | dices, body co<br>9, 1994, 470                  | mposition<br>479.        | n, and bor              | e turnos   | ver in healthy        |
|                          | G.S          | Johans<br>growtl  | sson et<br>n horme | al., <u>Trea</u><br>one in th | itment o              | of growth hormon<br>prospinal fluid and | e-deficient adults with r                        | ecombinan<br>rs Neuroer     | t human grov<br>docrinology,                    | vth hormor<br>Vol. 61, 1 | ne increas<br>995, 57-6 | ses the c  | oncentration of       |
|                          | G.T          |                   |                    |                               |                       |   | or diagnosing growth ho                          |                             |   |                          |                         |            | t 2, 1996, S61-       |
| EXAMINER                 | C            | l                 | 11                 | 7                             | /                     |   | DATE CONS  | SIDERED                     | 611   | 4/                       | 04                      |            |                       |
| EXAMINER:                | Initial if c | itation c         | onside:<br>munica  | red, whe                      | ther or               | not in conforman                        | ce. Draw line through c                          | itation only                | y if not in cor                                 | ıformance_               | anchot co               | nsidered   | I. Include a          |

Substitute Disclosure Form (PTO-1449)

Sheet 7 (G) of 12

|                     |        |  |       |        |               |                         |                    |                        |                              |                             | _  |                  |                     |            | She                | et 9(1) of        | 12          |
|---------------------|--------|--|-------|--------|---------------|-------------------------|--------------------|------------------------|------------------------------|-----------------------------|--|------------------|---------------------|------------|--------------------|-------------------|-------------|
| SUBSTITUTE I        | ORM PT | O-14   | 49    |        |               |                         | Į<br>P             | U.S. DEPA              | ARTMENT<br>AND TRAD          | OF COMMERCE<br>EMARK OFFICE | 3  | ATTY.            | DOCKET NO<br>1/04   | ).<br>     | SERIAL<br>09/939,9 |                   | E           |
|                     |        |  |       |        |               |                         |                    | OSURE<br>LICANT        |                              |                             |  | APPLIC<br>Michae | CANT<br>1 B. Foster |            |                    | 0,                |             |
|                     |        |  | (U:   | se se  | vera          | l sheet                 | s if ne            | cessary)               |                              |                             | FILING DATE August 27, 2001  GROUP APP       |                  |                     |            |                    | APR O             | 5 200       |
| (37 CFR 1.98(b)     | ))     |  |       |        |               |                         |                    |                        |                              |                             |  | August           | 27, 2001            | ·          | - 1000             | 10 16 2           | - (4)       |
|                     |        | _  |       |        |               |                         |                    |                        | U.S. PATE                    | NT DOCUMENTS                | <u> </u>                                     |                  |                     |            | FILING DATE        |                   |             |
| EXAMINER<br>INITIAL |        | L,   | PAT   | ΓEN    | T NI          | <b>ЈМВ</b> Е            | R                  | ISSUE PATENTEE         |                              |                             | <u>.                                    </u> |                  | CLASS               | SUBC       | CLASS              | IF APPROP         | AIE         |
|                     | I.A    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     | ļ          |                    |                   |             |
|                     | I.B    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     | <u> </u>   |                    |                   |             |
|                     | I.C    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.D    |  |       |        |               |                         |                    |                        |                              |                             |  |                  | <u> </u>            |            |                    |                   |             |
|                     | I.E    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     | ļ          |                    |                   |             |
|                     | I.F    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.G    |  |       |        |               |                         |                    |                        |                              | <u> </u>                    |  |                  |                     | ļ          |                    |                   |             |
|                     | I.H    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.I    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     | ļ          |                    |                   |             |
|                     | I.J    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     | <u> </u>   |                    |                   |             |
|                     | I.K    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     |        |  |       |        | F             | OREI                    | GN PA              | TENTS                  | OR PUBLIS                    | HED FOREIGN F               | 'ΑΊ  | TENT A           | PPLICATION          | S          |                    |                   |             |
|                     |        |  |       |        | MEN<br>BER    |                         | P                  | UBLICA<br>DATI         |                              | COUNTRY PATENT OF           |  |                  | CLASS               | SUBC       | LASS               | TRANSLA<br>(YES/N |             |
|                     | I.L    |  |       |        |               | -                       |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.M    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   | _           |
|                     | I.N    |  |       |        |               |                         |                    |                        |                              |                             |  |                  | ! <u></u>           |            |                    |                   |             |
|                     | I.O    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | 1.P    |  | -     |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.Q    |  |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     |        |  |       |        |               | OTHE                    | R DO               | CUMEN                  | TS (Includin                 | g Author, Title, D          | ate  | , Place o        | f Publication       | )          |                    | <u> </u>          |             |
| coc                 | I.R    | Dra<br>Bla   | ike a | and N  | Mons<br>Scien | on, <u>In</u><br>ce Ltd | dividu:<br>., Osne | al suscept<br>ey Mead, | ibility and C<br>Oxford, 199 | GH dosing regiments         | n in   | adults i         | n Growth Hor        | mone The   | rapy John          | P. Monson, Ed     | l. <b>,</b> |
|                     | I.S    | Nass et al., Effect of growth hormone (hGH) replacement therapy on physical work capacity and cardiac and pulmonary function in patients with hGH deficiency acquired in adulthood, Journal of Clinical Endocrinology and Metabolism, Vol. 80, No. 2, 1995, 552-557. |       |        |               |                         |                    |                        |                              |                             |  |                  |                     |            |                    |                   |             |
|                     | I.T    | Ro.  | sen ( | et al. | , <u>De</u>   | creased                 | i psyci            | hological              | well-being i                 | n adult patients w          | ith_   | growth           | normone defic       | eiencyClin | ical Endoc         | erinology, Vol.   | 40,         |
| EXAMINER            | Ce     |  | 2     | 4      | /             | /                       |                    |                        |                              | DATE CO                     | 6/14/04                                      |                  |                     |            |                    |                   |             |

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and ot considered copy of this form with next communication to applicant.

Sheet 10 (J) of 12 SERIAL NO. U.S. DEPARTMENT OF COMMERCE ATTY. DOCKET NO. SUBSTITUTE FORM PTO-1449 09/939,962 RENAS/04 PATENT AND TRADEMARK OFFICE (MODIFIED) APPLICANT INFORMATION DISCLOSURE Michael B. Foster STATEMENT BY APPLICANT **GROUP** FILING DATE (Use several sheets if necessary) 1463 August 27, 2001 (37 CFR 1.98(b)) U.S. PATENT DOCUMENTS **FILING DATE ISSUE EXAMINER** IF APPROPRIATE **SUBCLASS PATENTEE CLASS** PATENT NUMBER DATE INITIAL J.A J.B J.C J.D J.E J.F J.G J.H J.J J.K FOREIGN PATENTS OR PUBLISHED FOREIGN PATENT APPLICATIONS TRANSLATION **PUBLICATION COUNTRY OR DOCUMENT** (YES/NO) **SUBCLASS CLASS** PATENT OFFICE DATE NUMBER J.L J.M J.N J.O J.P J.O OTHER DOCUMENTS (Including Author, Title, Date, Place of Publication) Rosen et al., Cardiovascular risk factors in adult patients with growth hormone deficiency Acta Endocrinologica, Vol. 129, 1993, 195-J.R con Rosen and Bengtsson, Premature mortality due to cardiovascular disease in hypopituitarism The Lancet, Vol. 336, 1990, 285-288. J.S Rudman, M.D. et al., Effects of human growth hormone in men over 60 years old The New England Journal of Medicine, Vol. 323, No. J.T 1, 1990, 1-6.

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and considered. Include a copy of this form with next communication to applicant.

**EXAMINER** 

DATE CONSIDERED

Sheet 11 (K) of 12

| SUBSTITUTE<br>(MODIFIED) | FORM P       | TO-1449  | )                           |                                   |  | ENT OF COMMERCE RADEMARK OFFICE                    | ATTY.<br>RENAS                               | DOCKET NO<br>5/04              | O.<br>                 | SERIAL NO.<br>09/939,962 \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ \ |  |  |
|--------------------------|--------------|--|-----------------------------|-----------------------------------|--|--|--|--------------------------------|------------------------|--|--|--|
|                          |              |  |                             |                                   | SCLOSURE<br>APPLICANT                          |  | APPLIC<br>Michae                             | CANT<br>I B. Foster            |                        |  | 9 5 2004                               |  |
| (27.000.1.000            |              | (  | Use sev                     | eral sheets                       | if necessary)                                  |  | FILING                                       | DATE<br>27, 2001               |                        | GROU   | 16H                                    |  |
| (37 CFR 1.98(            | 0))          |  |                             |                                   | IIS PA   | TENT DOCUMENTS                                     | August                                       | 27,2001                        |                        | 2033   | TRADENA                                |  |
| EXAMINER                 | Τ            | 1  |                             |                                   | ISSUE  | TENT DOCOMENTS                                     | · · · · ·                                    |                                |                        | ·  | FILING DATE                            |  |
| INITIAL                  |              | P.A  | TENT                        | NUMBER                            | DATE   | PATENTEE   |  | CLASS                          | SUBCLASS               |  | IF APPROPRIATE                         |  |
|                          | K.A          |  | ╁╌┼╴                        | +++                               | <u> </u>                                       |  |  |                                |                        |  |  |  |
|                          | K.B          | +  | +-                          | ╂╌┼╌┼                             | <del>_</del>                                   |  |  | <u> </u>                       |                        |  | _                                      |  |
|                          | K.C          | ╁╂   | ++                          | +++                               |  |  |  |                                | -                      |  |  |  |
|                          | K.D          | ╁┼╌  | ++                          | ++                                |  |  | <u>.                                    </u> |                                |                        |  | ļ                                      |  |
|                          | K.E          | ┼-┼-   | ++                          |                                   |  |  |  | <u> </u>                       | <u> </u>               | <u>_</u> -   |  |  |
|                          | K.F          | <del>                                     </del> | ++                          | ╁┼┼┤                              |  | <del>-,-</del>                                     |  |                                | <u> </u>               |  |  |  |
|                          | K.G          |  | +                           | +                                 |  |  |  | <u> </u>                       |                        | <u> </u>   |  |  |
|                          | K.H          | ┼┼-  | ++                          | +++                               |  |  |  | <br>                           | <del> </del>           |  |  |  |
|                          | K.I          |  | ++                          | ╂╌╂╌╂                             |  | <del> </del>                                       |  | <u> </u>                       |                        |  | -                                      |  |
|                          | K.J          | ++   | ++                          |                                   |  | <del></del>  |  |                                | <del> </del>           | ···  |  |  |
| <u> </u>                 | K.K          |  |                             | FOREIGN                           | I PATENTS OR PLIE                              | LISHED FOREIGN PA                                  | TENT AP                                      | PLICATION:                     | <br>S                  |  | ــــــــــــــــــــــــــــــــــــــ |  |
|                          |              | DO   | OCUME                       | - T                               | PUBLICATION                                    | COUNTRY OF   | T  |                                |                        |  | TRANSLATION                            |  |
| <u>.</u>                 |              |  | NUMBE                       |                                   | DATE   | PATENT OFFIC                                       | CE   | CLASS                          | SUBCL                  | LASS   | (YES/NO)                               |  |
|                          | K.L          |  |                             |                                   |  |  |  |                                |                        |  |  |  |
|                          | K.M          |  |                             |                                   |  |  |  |                                |                        |  |  |  |
|                          | K.N          |  |                             |                                   |  |  |  |                                |                        |  |  |  |
|                          | K.O          |  |                             |                                   | ·  |  |  |                                | ··                     |  |  |  |
|                          | K.P          |  |                             |                                   |  |  |  |                                |                        |  |  |  |
|                          | K.Q          | <u> </u>   |                             |                                   |  |  |  |                                | _                      |  |  |  |
|                          | <del>_</del> |  |                             |                                   |  | ding Author, Title, Date                           |  |                                |                        |  |  |  |
| Con                      | K.R          | Russel<br>in adul                                | I-Jones<br>It growt         | et al., <u>The c</u><br>h hormone | effect of growth hormodeficient patients, Clir | one replacement on seru<br>nical Endocrinology, Vo | m lipids, l<br>1. 41, 199                    | ipoproteins, :<br>4, 345-350.  | apolipopro             | teins and  | cholesterol precursors                 |  |
|                          | K.S          | Salom<br>growth                                  | on et al.<br>hormo          | , <u>The effec</u><br>ne deficien | ts of treatment with recy, The New England     | combinant human growt<br>Journal of Medicine, Vo   | th hormon<br>ol. 321, No                     | e on body co<br>o. 26, 1989, 1 | mposition<br>797-1803. | and meta   | bolism in adults with                  |  |
|                          | К.Т          | Schwa<br>A, Vol                                  | rtz, <u>Tro</u><br>. 50A, 1 | pic factor s<br>995, 151-1        | upplementation: effec                          | et on the age-associated                           | changes ir                                   | 1 body compo                   | sition The             | Journal o  | of Gerontology, Series                 |  |
| EXAMINER                 | 0            |  | 10                          | /                                 |  | DATE CONSI   | IDERED                                       | 6                              | 5/19                   | 1/0  | 4                                      |  |
| EXAMINER:                | Initial if c | itation co                                       | onsidere                    | d, whether                        | or not in conformance                          | e. Draw line through cit                           | ation only                                   | if not in con                  | formance :             | anchot co  | nsidered. Include a                    |  |

Sheet 12 (L) of 12 SUBSTITUTE FORM PTO-1449 SERIAL NO. U.S. DEPARTMENT OF COMMERCE ATTY. DOCKET NO. 09/939,962 PATENT AND TRADEMARK OFFICE RENAS/04 (MODIFIED) APPLICANT INFORMATION DISCLOSURE Michael B. Foster STATEMENT BY APPLICANT **GROUP** FILING DATE (Use several sheets if necessary) 1655 August 27, 2001 (37 CFR 1.98(b)) **U.S. PATENT DOCUMENTS FILING DATE ISSUE EXAMINER** IF APPROPRIATE INITIAL PATENT NUMBER DATE **PATENTEE CLASS SUBCLASS** L.A L.B L.C L.D L.E L.F L.G L.H L.I L.J L.K FOREIGN PATENTS OR PUBLISHED FOREIGN PATENT APPLICATIONS TRANSLATION **PUBLICATION COUNTRY OR** DOCUMENT (YES/NO) **CLASS SUBCLASS** NUMBER DATE PATENT OFFICE L.L L.M L.N L.O L.P L.Q OTHER DOCUMENTS (Including Author, Title, Date, Place of Publication) CH Shore et al., Bone mineral status in growth hormone deficiency, J. Pediatrics, Vol. 96, No. 3, part 1, 1980, 393-396. L.R Sönksen and Weissberger, Growth hormone deficiency in adults Growth Genetics and Hormones, Vol. 14, No. 3, 1998, 41-48. L.S L.T

EXAMINER: Initial if citation considered, whether or not in conformance. Draw line through citation only if not in conformance and to considered. Include a copy of this form with next communication to applicant.

**EXAMINER** 

DATE CONSIDERED